

## CLAIMS

1/ Compound capable of cross-linking a stimulatory receptor with a KIR.

5

2/ Compound according to claim 1, characterized in that it is capable of specifically regulating the activation of a KIR.

10 3/ Compound according to claim 1 or 2, characterized in that it is capable of regulating the activation of a stimulatory receptor.

15 4/ Compound according to any of the preceding claims characterized in that said stimulatory receptor is an ITAM-bearing receptor such as KAR, Fc $\epsilon$ RI, CD3/TCR, CD16, receptors related to tyrosine kinase activities or a receptor sub-unit such as CD3 $\zeta$ , CD3 $\epsilon$ , CD3 $\gamma$ , CD3 $\delta$  or Fc $\sigma$ RIY.

20 5/ Compound according to any one of claims 1-4, characterized in that said KIR is a IgSF member, particularly selected from the group comprising CD158, CDw159, CDw160, or said KIR is lectin-like, such as the CD94-NKG2A/ B heterodimer.

25

6/ Compound according to any of the preceding claims, characterized in that said KIR is expressed on a NK cell, on a T cell, on a mast cell or on a monocyte or is recombinantly expressed.

30

7/ Compound according to any of the preceding claims, characterized in that it is capable of inducing the regulation of free calcium concentration in a cell, particularly of inducing the regulation of calcium influx into a cell and/or of inducing the regulation of calcium mobilization from intracellular compartments.

8/ Compound according to any of the preceding claims, characterized in that it is capable of inducing the recruitment by said KIR or KIR homologue of a phosphatase  
5 selected from the group consisting of SHP-1, SHP-2.

9/ Compound according to any of the preceding claims, characterized in that it is essentially a polypeptide, a glycoprotein or a carbohydrate.

10 10/ Compound according to any of the preceding claims, characterized in that said compound is a bispecific reagent and/or a chemical inducer of dimerization.

15 11/ Compound according to any of the preceding claims, characterized in that said compound is a bispecific antibody, comprising at least one Fab, Fd, Fv, dAb, CDR, F(ab')<sub>2</sub>, VH, VL, ScFv fragment.

20 12/ Compound according to any of the preceding claims, characterized in that it is capable of cross-linking said KIR with said stimulatory receptor in the extracellular domain of a cell.

25 13/ Compound according to any of the preceding claims, characterized in that it is capable of crossing through a lipid bi-layer, and is liposoluble and/or associated with a drug-delivery system.

30 14/ Compound according to any of the preceding claims, characterized in that it is capable of cross-linking said KIR with said stimulatory receptor in the intracellular domain of a cell.

35 15/ Compound according to any of the preceding claims, characterized in that it is capable of modulating the

release of serotonin and/or of inflammatory mediators by a cell expressing Fc $\epsilon$ RI, such as a mast cell, and/ or of modulating cytokine release, such as Interleukin-6, Tumor Necrosis Factor Alpha release, from a cell such as a mast cell or a NK cell and/or of modulating interleukin production such as the IL-2 production and/or the  $\gamma$ -interferon production from a peripheral blood cell and/or of modulating the proliferation of peripheral blood cells.

10 16/ Compound according to any of the preceding claims, characterized in that it is capable of controlling the host tolerance to allogeneic grafts and/or the graft toxicity against a host tissue.

15 17/ Nucleic acid coding for a polypeptide according to any one of claims 9-16.

18/ Cell transfected by a nucleic acid according to claim 17.

20 19/ Pharmaceutical preparation comprising a compound according to any one of claims 1-16 or a nucleic acid according to claim 17 or a cell according to claim 18 in a physiologically acceptable vehicle, in a therapeutically-effective amount useful for modulating an animal cell function involved in a disease selected from the group consisting of immunoproliferative diseases, immunodeficiency diseases, cancers, autoimmune diseases, infectious diseases, viral diseases, inflammatory responses, allergic responses or involved in organ transplant tolerance.

25 30 35 20/ Method for the *in vitro* or *ex vivo* diagnosis of a cell disregulation, comprising the step of estimating of the relative proportion of co-aggregated KIR vs. non-co-aggregated KIR by:

- contacting a biological sample with a compound according to any one of claims 1-16 or with a nucleic acid according to claim 17 or a cell according to claim 18, and
- revealing the reaction product possibly formed.